

SEQ. ID. NO.:1

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TGAGCCTATTGGCCCCACAGCTCTCAGGAGCAAGAGTGATGTACAGGACATTGTGAGCAAGAAGTGGGT
GCTGCAAACTGCATAACCCCCCTCTACCGGCAAGACACCGAGTGCTCACACAGAGCTTACTCGTAGGAC
TTGCCAGCTGGTTAAGACACACCCTGCCATTTTCTCTAACAAGCAGGAGTTCAGTTCAGTTCACAGGGAT
GGGGTGGGACCAGGATGGCCACTTTGATCACATGGGAGGGGCGTGGTGTGTGTCAGTTAGGAACAAAGTC
TCCCCCTATTTAAGTCCAGCGCTCTGTGCTTTAGTTGATCCCTGGTGTCTCGTGTCTTTGTCTGCTGCTG
TCCCGCCACCAGCCCCAGCCATGCAGGGACCCTGGGTGCTGCTGCTGCTGGGCCTCAGGCTACAGCTGTC
CCTTAGTGTCATTCCAGGTAATGAGGCTCCTTCCAATGAACACCCCATTTCCACCCATGGACCCCTTCATG
CTGACCCTTCCTCTGTCTATTCCCTTGGCCAGTGGAGGAGGAGAACCCGGCCTTCTGGAACAAGAAGGCAG
CCGAGGCCCTGGATGCTGCCAAGAAGCTGCAGGCCATTTCAGACATCAGCTAAGAACCTCATCATCTTCCT
GGGTGACCGTGAGTGTGTGAGCGAGGCCTGCCACCCTGGGGCCCTTGTAATCCAAGTACCCAGGGCCACT
GGTGGGTACGGACAGGCCTCAGGGTTCAGTCTGACGAGGTTCTGCTCCTTCAGGAATGGGGGTACCAAC
AGTGACAGCCACCAGGATCCTAAAGGGACAGTTGGAAGGTCATCTAGGACCTGAGACACCCCTAGCCATG
GACCGCTTCCCATATATGGCTCTGTCCAAGGTGAGTTCCTTAGCCACATCTGAAATGACTGATGGGATCCA
GGGCAAGGGAGGCAGAGAGGCTCGGGTGAAGAAATAAATGTCTGCTTTGAGCCAGTTGGGGTGTCTCTG
TCCCCAGACATACAGTGTGGACAGACAGGTTCCAGACAGTGCAAGCACGGCCACCGCCTACCTGTGTGGG
GTCAAGACCAACTACAAGACCATCGGCTTGAGTGACGCCGCGAGATTTCAGGAGTGCAACACCACATTTG
GCAATGAGGTCTTCTCAGTGATGTACCGTGCCAAGAAAGCAGGTGAGTTGGAGCCAGGCTCAGCTATGGG
GGGCAAGCCTAGGGGACTGGATGTCTCACCCCTGACCTTTGCCGTCTTCAGGAAAATCCGTAGGTGTGGTG
ACCACCACCAGAGTGACAGCACGCTCTCCCTCGGGCACATATGTTACACAGTGAACCGCAATTGGTATG
GGGATGCTGACATGCCTGCCTCTGCGCTGCGGGAAGGTTGCAAGGACATTGCTACACAATCATCTCCAA
CATGGACATTAATGTAAGGATAAGCATGTCAAAGGGAGAGGGTAAGGGGAGGGAGAGGAGGAGAAGGAGG
GGGAGGGAGGGGAGGTGAGGGGGTCAAGGGGGGAAGGGGTGGTCCCAGGCAAACCTTGTAGACTGAAC
TCCCTGGATCTTCTGGGGTCTTTGAGGGCCGGGTAGTTTCAGTTCCCACATACCTGGTGAGGAGCTAGGGA
CTGGCAGGAAAAGGAGGCAGAAGACAACCTAAAGTTCACCTTCCTTCATCCTCTCTGACCACAGGTGATC
CTTGGTGGGGGGCGAAAATACATGTTTCCCTGCTGGAACCCACAGCCCCAGATATCCAAATGATGCTAATG
AGACTGGAACCAGATTGGATGGCAGGAATCTGGTGCAAGGAATGGCTGTCAAAGCACCAGGTGACCGACTG
CAGAATATTAGTGATACAGTGGAGACCAGGGAAGGGCTTTGAACCTTACCAGTTGCTTATGTCCCTCTAG
GGATCCCAGTATGTTTGGAAATCGTGAACAATCATTCAGAAGGCCAGGATCCGTGAGTGACATACCTCA
TGGGTAATGGCCCCACACTTCTGCACTGGTACACCTCACATGGCAACCACTGATCCTCTGTGTATATAT
GTACCGTGACCCCACTGCCAAGCTTGGTGGTCAACAGTATATATTTTGGTTTTGTACCTCAGGCCTCTTT
GAGCCTGTAGACACAAAATTTGATATTCAACGAGATCCCCTGATGGACCCATCTCTGAAGGATATGACAG
AGACGGCCGTGAAAGTGCTAAGCAGGAACCCCAAAGGCTTTTATCTCTTTGTGGAGGGTGAGTCTCCAAG
CTCCCATGGAAAGAGGGGACAATGGACAGGGACAGGCTCAAGCTCACTGGCTTCCTGTCAGGGGGCCGAAT
CGACCGTGGTCAACATCTGGGCACAGCTTATCTGGCGCTGACTGAGGCTGTGATGTTGACTTAGCCATC
GAGAGGGCCAGCCAGCTCACTAGTGAACGCGACACTCTGACCATAGTCACTGCTGACCACTCCCATGTCT
TCTCCTTTTGGTGGCTACACACTTCGAGGGACCTCCATCTTCGGTAGGTTCCGGAACAGTGGCAGGCTGTC
AATTACGTACAGAATACTTCTGAGCCATCGTTTCTCTGTCTGTAAAATGGACAGAAATGGCACCTGCCT
TGTGGGGATCTAGCAACGACTGAACCACTGGCCAGGCAAAAGGCGGGGGCTCGTCTAAGCATCATTTCTTG
GCAGGAAAAGTGTCCCTCTTCCCCATGCAGGGCTGGCTCCCCTCAATGCTCTGGACGGCAAGCCCTAC
ACCTCCATCCTGTATGGCAACGGCCCAGGCTATGTGCGTACAGGGGAAAGACCCAACGTACCGCCGCTG
AAAGCAGTGAGTGCGGTGGGGTGGCTTGCCCTGAAGGTCGGGTAGAGGTGACTCAGATCAGAGTCTCTCC
CTTAACATCTTGTCCCTACCAGGTGGCTCATCGTACCGCAGGCAGGCTGCTGTGCCGGTGAAGTCGGAGA
CCCACGGCGGGGAGGACGTGGCGATATTGCGCGTGGCCCCGAGGCGCACTTGGTGCACGGGGTGCAGGA
GCAGAATACTACGCGCACGTGATGGCCTCTGCAGGCTGCCCTGGAGCCCTACACCGACTGCGGCTTGGA
CCCCCTGCAGATGAAAGCCAGACCACACGACAACCCGCCAGACCACCATCACCACCACCACCACCACCA
CCACCACCACAACCACCCCGGTCCATAACAGCGCCAGAAGCCTGGGGCCAGCCACCGCCCCGCTGGCTCT
GGCGCTGCTGGCCGAATGCTGATGCTACTACTAGGGGCTCCTGCGGAGTCTAACTCCAGCACATCTA
GGCTCCACCCACTAGGTCCCACGCCCTCACCTGGTCCTTCCCTTCCCTGACCTCAGTGCTCCCTGCATTC

FIGURE 1A

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TCCCTGCGGGCTCTACCCCAGGATCCTCTCTCTGTCTTTCTGCTACTGGCCTCATGTCTAGCCCTACCTT
GCATTGCAGCTTCCAGGTTCCCTCCTACCCAGGCACTCACAAAGGCCAATCACCTCTGAGCTAGCAGCCAG
CCTCAGACCCACAGAGTTACTTCTCCCCAGGCAGCATGACCACCAAGGCCCTGGACCTCCCGGGGCAAT
CCGGACTCTCCTTTTGGCCCTCATCCATCAGCCCCCTAGAAAAAGATAGGATCCCGCAATAATTTGTGGAGG
ACCAAACATGCACCTGCCCATTGGCACTTCTCCGAGCTTGAATCCATCTTACAGGCTCTGTACCCAGGA
CTAAGGCACAAGAGAACACAGAGAGAGGCTGTCTTCCCACTACTCCTCGGTCTAATCTGCTGGCAGGTGG
CAAGGCTACGGTGCTGGGTACCCCTAGCCAGCCTTTGACATAGTTCTTCCCTCGATGTCTCTGGACCAGCTC
CACATTCAAAACCATCATGGCTCAGCCATACCAACCCACAGAGCGAAGATTCTGAAATCGTTCAGCCCTT
TCATGTCTATTGCCCAGCTAGGAGATTCAAAGAGCTGTACCCACCCCACTCTCAGGTCATCTCAGGTTG
CACCTAAATTTCTGAACTGAGAAAAGTCCCTAACTTCCCAGGTCTGCATTCCCCTGGGGAGAGTCAAGTC
AATAATAAAAAGATGTATTCAATACAATAGCAATAGTCATTTTCTTTTCTTTCGGCTCAAAACCAGAGCC
TAGTGCTGTCTAGGAACGTGCTCTGCCACTGATCCATAGCCCCATATCATCTCCTCCCCCTCCCCCTCTCCT
CCTCCCTCTTCTCCTTCCCCCTCCTCCTCTATGACTCTGTAGCCCAAGCTGGCCTCAAATTTATGACAGT
CCACTTGCTACAGTCTCCCAGATGCTGGATTTTAAAGTGTGAGCCACACTCCTAGCATCTTAGTAGGACCT
TTGCAGAAGGAAAGCCTGAAGTGCTGGAGCACTGAGTTCAGATGGGGGAGGGTAATAGTGGAGCCTCA
GTTGGAGAGAGACAGCCAGCTGAGCAAGATCCTGAATGAGGTGAAGGCCTGAGCCAACACCACACAGCAG
TGCTAATCCCCCACCCTCCAGGCCAGCGATCAGCTGGAAGGTTGCAACGACTGGGTGAGAGAGGGTGGCT
GGGACAGAGGATGCAAAGCTGGAGCTGCAAGGAGCTGTGGGAGGAGAGGAAGAAGCTTTAAAATCCATGGC
AGTGTGGTCACAAGCCTTTGAATAAGAATTCAGGACGTGGTACTTTTTCTATTGCAGGAAATATGCAATC
TTTTCCCTTTTTTCTGTTTTTTTTTCCATGGGGGGTGGGAATGGGTGTTAGATATAGGAGCTGGTCA
GCCAGAGGGGAGATGCAGACCCTAACCATCTCTGACTTGCAATTGGAAGTGGTGGAGCACCACCCAGTA
TAGTTCTTGGCCCTGTCTAACCTGCCAATGAGGACATTTGAAGGAATTACGTAAAGGTGGATTAAAGCT
GTGTTTCTCAGTAAGTTTTGCAACACTACAAATTTATCTGTACATTTATGAAGGTACAAAACACACTTT
GCTCCCACTAGTAATATTAGGAAGATTGAATATGCATCCTTATTTGCTAAAATCTTGATTTAACACTGTG
AAACATCAATTTCGAAATCTTGGCTCTCGGAGTAGTTTATTTCAATTCCGGATTTTAGTGGCTGTGAGAA
AATATGGGAGCTGAATGGAAAAAGGCCATCGTTAACAAAGCTT

SEQ. ID. NO.:2

MQGPWVLLLLGLRLQLSLSVIPVEENPAFWNKAAEALDAAKKLQPIQTSAKNLIIFLGDGMGVPTVTAT
RILKGQLEGLGPETPLAMDRFPYMALSKTYSVDRQVPDSASTATAYLCGVKTNKYKTIGLSAAARFDQCNT
TFGNEVFSVMYRAKKAGKSVGVVTTTRVQHASPSGTIVHTVNRNHWYGDADMPASALREGCKDIATQLISNM
DINVILGGGRKYMFPAGTPDPEYPNDANETGTRLDGRNLVQEWLSKHQGSQYVWNREQLIQKAQDPSTVYL
MGLFEPVDTKFQDIQRDPLMDPSLKDMTETAVKLSRNPKEFYLFVEGGRIDRGHHLGTAYLALTEAVMFDL
AIERASQLTSERDTLTIVTADHSHVFSFGGYTLRGTSIFGLAPLNALDGKPYTSILYGNPGPYVGTGERPN
VTAAESSGSSYRRQAAVPVKSETHGGEDVAIFARGPQAHVHGVQEQNYIAHVMAAGCLEPYTDCGLAPP
ADESQTTTTTRQTTITTTTTTTTTTTTTTPVHNSARSLGPATAPLALALLAGMLMLLLGAPAES

FIGURE 1B

Underlined = deleted in targeting construct

Bold = sequence flanking Neo insert in targeting construct

AAGCTTAATTGGGGGCCAAGTAGACAGCAGGACATTCAAGTGTGCCTTGTTCCTTTGTCT
TTTGGCTCCAGGTATCAGCAAGCCAAACAAAGGCCCTCATCTAAGCTGTGTTCTTCAGG
CCTACCTCCAGCGCCAGAATGAGCCTATTGGCCCCACAGCTCTCAGGAGCAAGAGTGA
TGTACAGGACATTGTGAGCAAGAAGTGGGTGCTGCAAACATGCATAACCCCCCTCCTACCG
GCAAGACACCGAGTGTCTCACACAGAGCTTACTCGTAGGACTTGCCAGCTGGTTAAGACAC
ACCTTGCCATTCTCTTAACAAGCAGGAGTTCAAGTTCAGTTCACAGGGATGGGGTGGGAC
CAGGATGGCCACTTTGATCACATGGGAGGGGCGTGGTGTGTGTCAGTTAGGAACAAAGTC
TCCCCCTATTTAAGTCCAGCGCTCTGTGCTTTAGTTGATCCCTGGTGTCTCGTGTCTTTG
TCTGCTGCTGTCCCGCCACCAGCCCCAGCCATGCAGGGACCCCTGGGTGCTGCTGCTGCTG
GGCCTCAGGCTACAGCTGTCCCTTAGTGTCAATCCAGGTAATGAGGCTCCTTCCAATGAA
CACCCCATTTCCACCCATGGACCTTTCATGCTGACCTTCCCTCTGCTATTCCCTTGGCCA
GTGGAGGAGGAGAAACCCGGCCTTCTGGAACAAGAAGGCAGCCGAGGCCCTGGATGCTGCC
AAGAAGCTGCAGGCCATTGATCAGACATCAGCTAAGAACCTCATCATCTTCTTGGGTGACGGT
GAGTGTGTGAGCGAGGCCTGCCACCTGGGGCCCTTGTACTCCAAGTACCCAGGGCCACT
GGTGGGTACGGACAGGCCTCAGGGTTCAAGTCTGACGAGGTTCTGCTCCTTCAGGAATGG
GGGTACCAACAGTGACAGCCACCAGGATCCTAAAGGGACAGTTGGAAGGTCATCTAGGAC
CTGAGACACCCCTAGCCATGGACCGCTTCCCATATATGGCTCTGTCCAAGGTGAGTCTT
AGCCACATCTGAAATGACTGATGGGATCCAGGGCAAGGGAGGCAGAGAGGCTCGGGTGAA
GAAATAAATGTCTGCTTTGAGCCAGTTGGGGTGTCTCTGTCCCCAGACATACAGTGTGG
ACAGACAGGTTCCAGACAGTGCAAGCACGGCCACCGCCTACCTGTGTGGGGTCAAGACCA
ACTACAAGACCATCGGCTTGAGTGCAGCCGCGAGATTCGACCAAGTGAACACCACATTTG
GCAATGAGGTCTTCTCAGTGATGTACCGTGCCAAGAAAGCAGGTGAGTTGGAGCCAGGCT
CAGCTATGGGGGGCAAGCCTAGGGGACTGGATGTCTACCCCTGACCTTTGCGGCTTTCAG
GAAATCCGTAGGTGTGGTGACCAACAGAGTGCAGCACGCTCTCCCTCGGGACAT
ATGTTTACACAGTGAACCGCAATTGGTATGGGGATGCTGACATGCCTGCCTCTGCGCTGC
GGGAAGGTTGCAAGGACATTGCTACACAACCTCATCTCCAACATGGACATTAATGTAAGGA
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TCCCTGGATCTTCTGGGGTCTTTGAGGGCCGGGTAGTTCAAGTTCACACATACCTGGTGAG
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CTCTCTGACCACAGGTGATCCTTGGTGGGGGGCGAAAATACATGTTTCTGCTGGAACCC
CAGACCCCGAGTATCCAAATGATGCTAATGAGACTGGAACCAGATTGGATGGCAGGAATC
TGGTGCAGGAATGGCTGTCAAAGCACCAGGTGACCGACTGCAGAATATTAGTGATACAGT
GGAGACCAGGGAAGGGCTTTGAACCTTACCAGTTGCTTATGTCCCTCTAGGGATCCCAGT
ATGTTTGAATCGTGAACAACCTCATTCAGAAGGCCAGGATCCGTCAGTGACATACCTCA
TGGGTAATGGCCCCACACTTCCCTGCCTGAGTACCTCAGATGGCAACCACTGATCCTCT
GTGTATATATGTACCGTGACCCCACTGCCAAGCTTGGTGGTCAACAGTATATATTTTGGT
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TGATGGACCCATCTCTGAAGGATATGACAGAGACGGCCGTGAAAGTGCTAAGCAGGAACC
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AATGGACAGGGACAGGCTCAAGCTCACTGGCTTCCCTGCAGGGGGCCGAATCGACCTGGT
CACCATCTGGGCACAGCTTATCTGGCGCTGACTGAGGCTGTGATGTTCCGACTTAGCCATC
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AGTCCCTCTCCCTTAACATCTTGTCCCTACCAGGTGGCTCATCGTACCGCAGGCAGGCTGC
TGTGCCGGTGAAGTCGGAGACCCACGGCGGGGAGGACGTGGCGATATTCGCGCGTGGCCC
GCAGGCGCACTTGGTGCACGGGGTGCAGGAGCAGAATACATCGCGCACGTATGGCCTC
TGCAGGCTGCCTGGAGCCCTACACCGACTGCGGCTTGGCACCCCTGCAGATGAAAGCCA
GACCACCACGACAACCCGCCAGACCACCATCACCACCACCACCACCACCACCACCACCAC
AACCACCCCGGTCCATAACAGCGCCAGAAGCTGGGGCCAGCCACCGCCCGCTGGCTCT

FIGURE 2A

GGCGCTGCTGGCCGGAATGCTGATGCTACTACTAGGGGCTCCTGCGGAGTCCTAAACTCC
AGCACATCTAGGCTCCACCCACTAGGTCCCACGCCCTCACCTGGTCCTTCCCTTCCCTGA
CCTCAGTGCTCCCTGCATTCTCCCTGCGGGCTCTACCCCAGGATCCTCTCTCTGTCTTTC
TGCTACTGGCCTCATGTCTAGCCCTACCTTGCAATTGCAGCTTCCAGGTTCCCTCCTACCCA
GGCACTCACAAAGGCCAATCACCTCTGAGCTAGCAGCCAGCCTCAGACCCACAGAGTTA
CTTCTCCCCAGGCAGCATGACCACCAAGGCCTTGGACCTCCCGGGGCAATCCGGACTCTC
CTTTTGCCCTCATCCATCAGCCCCCTAGAAAAAGATAGGATCCCGCAATAAATTTGTGGAGG
ACCAAACATGCACCTGCCCATTGGCACTTCCCTCCGAGCTTGAATCCATCTTACAGGCTCT
GTACCCAGGACTAAGGCACAAGAGAACACAGAGAGAGGCTGTCTTCCCCTACTCCTCGG
TCTAATCTGCTGGCAGGTGGCAAGGCTACGGTGTGGGTACCCTAGCCAGCCTTTGACAT
AGTTCTTCCCTCGATGTCTCTGGACCAGCTCCACATTCAAACCATCATGGCTCAGCCATA
CCAACCCACAGAGCGAAGATTCTGAAATCGTTTCAGCCCTTTCATGTCTATTGCCAGCTA
GGAGATTCAAAGAGCTGTACCCCACCCCACTCTCAGGTATCTCAGGTTGCACCTAAATT
TCTGAAGTGAAGAAAGTCCCTAAGTTCCTCAGGCTGTGATTCCCTGGGGAGAGTCAAGTC
AATAATAAAAGAATGTATTCAATACAATAGCAATAGTCATTTTCTTTTCTTCCGGCTCAA
AACCAGAGCCTAGTGCTGCTAGGAACGTGCTCTGCCACTGATCCATAGCCCCATATCAT
CTCCTCCCCCTCCCCCTCCTCCTCCTCCTCCTCCTCCTCCTCCTCCTCCTCCTCCTCCT
AGCCCAAGCTGGCCTCAAATTTATGACAGTCCACTTGCTACAGTCTCCCAGATGCTGGAT
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GTGTCTGGAGCACTGAGTTCAGATGGGGGAGGGGTAATAGTGGAGCCTCAGTTGGAGAGA
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TGCTAATCCCCCACCCCCCAGGCCAGCGATCAGCTGGAAGGTTGCAACGACTGGGTCAGA
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AGAACTTTAAATCCATGGCAGTGTGGTCACAAGCCTTTGAATAAGAATTCAGGACGTGG
TACTTTTTCTATTGCAGGAAATATGCAATCTTTTCCCCTTTTTTCTGTTTTTTTTTTCC
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GTGTTTCTCAGTAAGTTTGAACACTACAAATTTATCTGTACATTTATGAAGGTACAAA
AACACACTTTGTCTCCCACTAGTAATATTAGGAAGATTGAATATGCATCCTTATTTGCTAA
AATCTTGATTTAACACTGTGAAACATCAATTCGAAATCTTGGCTCTCGGAGTAGTTTATT
TCAATCCGGATTTTAGTGGCTGTGAGAAAATATGGGAGCTGAATGGAAAAGGCCATC
GTTAACAAAGCTT

FIGURE 2B

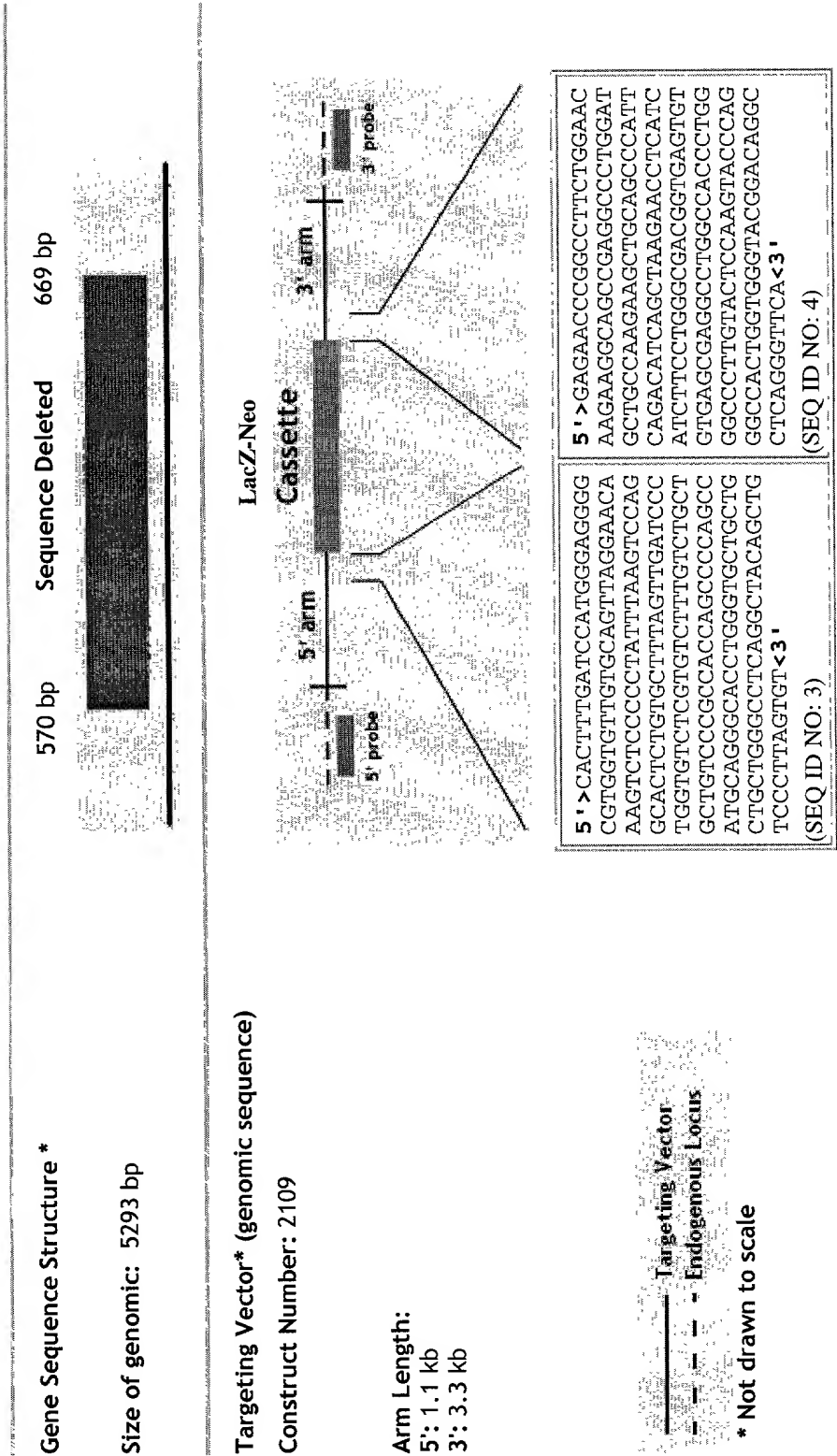


FIGURE 2C